NEWS HOURS

```
Welcome to STN International! Enter x:x
LOGINID:ssspta1202txn
PASSWORD:
TERMINAL (ENTER 1, 2, 3, OR ?):2
                     Welcome to STN International
NEWS
                 Web Page URLs for STN Seminar Schedule - N. America
NEWS
         Apr 08
                 "Ask CAS" for self-help around the clock
NEWS
         Apr 09
                 BEILSTEIN: Reload and Implementation of a New Subject Area
NEWS
         Apr 09
                 ZDB will be removed from STN
         Apr 19 US Patent Applications available in IFICDB, IFIPAT, and IFIUDB
NEWS
         Apr 22 Records from IP.com available in CAPLUS, HCAPLUS, and ZCAPLUS
NEWS
         Apr 22 BIOSIS Gene Names now available in TOXCENTER
NEWS
NEWS 8
         Apr 22 Federal Research in Progress (FEDRIP) now available
NEWS 9
         Jun 03
                 New e-mail delivery for search results now available
NEWS 10
         Jun 10
                 MEDLINE Reload
NEWS 11
         Jun 10
                 PCTFULL has been reloaded
NEWS 12
         Jul 02
                 FOREGE no longer contains STANDARDS file segment
NEWS 13
         Jul 22 USAN to be reloaded July 28, 2002;
                 saved answer sets no longer valid
NEWS 14 Jul 29
                 Enhanced polymer searching in REGISTRY
NEWS 15 Jul 30 NETFIRST to be removed from STN
NEWS 16 Aug 08
                 CANCERLIT reload
NEWS 17
         Aug 08
                 PHARMAMarketLetter(PHARMAML) - new on STN
NEWS 18
         Aug 08
                 NTIS has been reloaded and enhanced
NEWS 19
         Aug 19
                 Aquatic Toxicity Information Retrieval (AQUIRE)
                 now available on STN
NEWS 20 Aug 19
                 IFIPAT, IFICDB, and IFIUDB have been reloaded
NEWS 21
         Aug 19
                 The MEDLINE file segment of TOXCENTER has been reloaded
NEWS 22 Aug 26 Sequence searching in REGISTRY enhanced
NEWS 23 Sep 03
                 JAPIO has been reloaded and enhanced
NEWS 24 Sep 16 Experimental properties added to the REGISTRY file
NEWS 25 Sep 16 CA Section Thesaurus available in CAPLUS and CA
NEWS 26 Oct 01 CASREACT Enriched with Reactions from 1907 to 1985
NEWS 27 Oct 21 EVENTLINE has been reloaded
NEWS 28 Oct 24 BEILSTEIN adds new search fields
NEWS 29 Oct 24 Nutraceuticals International (NUTRACEUT) now available on STN
NEWS 30 Oct 25 MEDLINE SDI run of October 8, 2002
NEWS 31 Nov 18 DKILIT has been renamed APOLLIT
NEWS 32 Nov 25 More calculated properties added to REGISTRY
NEWS 33 Dec 02 TIBKAT will be removed from STN
NEWS 34 Dec 04 CSA files on STN
NEWS 35 Dec 17 PCTFULL now covers WP/PCT Applications from 1978 to date
NEWS 36 Dec 17 TOXCENTER enhanced with additional content
NEWS 37 Dec 17 Adis Clinical Trials Insight now available on STN
NEWS 38 Dec 30 ISMEC no longer available
NEWS 39 Jan 13 Indexing added to some pre-1967 records in CA/CAPLUS
NEWS 40 Jan 21 NUTRACEUT offering one free connect hour in February 2003
NEWS 41
         Jan 21 PHARMAML offering one free connect hour in February 2003
NEWS 42 Jan 29
                 Simultaneous left and right truncation added to COMPENDEX,
                 ENERGY, INSPEC
NEWS EXPRESS
              January 6 CURRENT WINDOWS VERSION IS V6.01a,
              CURRENT MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP),
              AND CURRENT DISCOVER FILE IS DATED 01 OCTOBER 2002
```

STN Operating Hours Plus Help Desk Availability

r

NEWS INTER General Internet Information NEWS LOGIN Welcome Banner and News Items

NEWS PHONE Direct Dial and Telecommunication Network Access to STN

NEWS WWW CAS World Wide Web Site (general information)

Enter NEWS followed by the item number or name to see news on that specific topic.

All use of STN is subject to the provisions of the STN Customer agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may result in loss of user privileges and other penalties.

FILE 'HOME' ENTERED AT 14:53:34 ON 30 JAN 2003

=> file reg
COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 14:53:42 ON 30 JAN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 28 JAN 2003 HIGHEST RN 482573-45-5 DICTIONARY FILE UPDATES: 28 JAN 2003 HIGHEST RN 482573-45-5

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

Uploading 09910702.str

L1 STRUCTURE UPLOADED

=> d l1

L1 HAS NO ANSWERS

L1 STR

Jecher Species

Structure attributes must be viewed using STN Express query preparation.

=> s 11

SAMPLE SEARCH INITIATED 14:54:39 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED - 0 TO ITERATE

100.0% PROCESSED 0 ITERATIONS 0 ANSWERS

1 ANSWERS

SEARCH TIME: 00.00.07

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

0 PROJECTED ITERATIONS: O TO 0 TO

PROJECTED ANSWERS:

0 SEA SSS SAM L1 L2

=> s l1 ful

FULL SEARCH INITIATED 14:54:55 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED - 5 TO ITERATE

100.0% PROCESSED SEARCH TIME: 00.00.01

L3 1 SEA SSS FUL L1

=> file marpat

COST IN U.S. DOLLARS SINCE FILE TOTAL

ENTRY SESSION FULL ESTIMATED COST 148.55 148.76

FILE 'MARPAT' ENTERED AT 14:55:09 ON 30 JAN 2003

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

5 ITERATIONS

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2003 American Chemical Society (ACS)

FILE CONTENT: 1988-PRESENT (VOL 104 ISS 15-VOL 138 ISS 4) (20030124/ED)

MOST RECENT CITATIONS FOR PATENTS FROM FIVE MAJOR ISSUING AGENCIES (COVERAGE TO THESE DATES IS NOT COMPLETE):

US 6500220 31 DEC 2002

20211496 19 NOV 2002 DE

1269992 01 JAN 2003

JP 2002373833 26 DEC 2002

WO 2003000260 03 JAN 2003

Structure search limits have been raised. See HELP SLIMIT for the new,

higher limits.

=> s l1 ful

FULL SEARCH INITIATED 14:55:17 FILE 'MARPAT'
FULL SCREEN SEARCH COMPLETED - 2248 TO ITERATE

99.6% PROCESSED 2238 ITERATIONS 1 ANSWERS

100.0% PROCESSED 2248 ITERATIONS 1 ANSWERS

SEARCH TIME: 00.00.36

L4 1 SEA SSS FUL L1

=> file caplus

COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST 104.95 253.71

FILE 'CAPLUS' ENTERED AT 14:56:06 ON 30 JAN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 30 Jan 2003 VOL 138 ISS 5 FILE LAST UPDATED: 29 Jan 2003 (20030129/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 14

L5 1 L4

=> d l5 1- ibib abs hitstr

YOU HAVE REQUESTED DATA FROM 1 ANSWERS - CONTINUE? Y/(N):y

L5 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 2002:90022 CAPLUS

DOCUMENT NUMBER: 136:129056

TITLE: .alpha.-Amino acid derivatives for inhibitors of

leukocyte adhesion mediated by VLA-4

INVENTOR(S): Konradi, Andrei W.; Pleiss, Michael A.; Thorsett,

Eugene D.; Ashwell, Susan; Welmaker, Gregory S.;

Kreft, Anthony; Sarantakis, Dimitrios; Dressen, Darren

B.; Grant, Francine S.; Semko, Christopher; Xu,

Ying-Zi

PATENT ASSIGNEE(S): Elan Pharmaceuticals, Inc., USA; American Home

Products Corporation

SOURCE: PCT Int. Appl., 125 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1 PATENT INFORMATION:

```
Regard
```

```
KIND DATE
      PATENT NO.
                                                      APPLICATION NO. DATE
      ______
                                                       -----
                                   20020131
      WO 2002008202
                           A2
                                                     WO 2001-US23075 20010720
           W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
           W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
      US 2002052470
                         A1 20020502
                                                    US 2001-910702 20010720
                                                   US 2000-220132P P 20000721
PRIORITY APPLN. INFO.:
OTHER SOURCE(S):
                               MARPAT 136:129056
      Disclosed are certain .alpha.-amino acid compds. which bind VLA-4.
      Certain of these compds. also inhibit leukocyte adhesion and, in
      particular, leukocyte adhesion mediated by VLA-4. Such compds. are useful
      in the treatment of inflammatory diseases in a mammalian patient, e.g.,
      human, such as asthma, Alzheimer's disease, atherosclerosis, AIDS
      dementia, diabetes, inflammatory bowel disease, rheumatoid arthritis,
      tissue transplantation, tumor metastasis and myocardial ischemia. The
      compds. can also be administered for the treatment of inflammatory brain
      diseases such as multiple sclerosis. Prepn. of N-[5-(2,2,2-
      trifluoroethyl)pyrimidin-4-yl]-L-4'-(1-methyl-4-methoxy-2-pyridon-3-
      yl)phenylalanine is described.
=> d his
       (FILE 'HOME' ENTERED AT 14:53:34 ON 30 JAN 2003)
      FILE 'REGISTRY' ENTERED AT 14:53:42 ON 30 JAN 2003
T.1
                   STRUCTURE UPLOADED
T<sub>1</sub>2
                 0 S L1
L3
                  1 S L1 FUL
      FILE 'MARPAT' ENTERED AT 14:55:09 ON 30 JAN 2003
L4
                  1 S L1 FUL
      FILE 'CAPLUS' ENTERED AT 14:56:06 ON 30 JAN 2003
L5
                 1 S L4
=> log y
COST IN U.S. DOLLARS
                                                               SINCE FILE
                                                                                   TOTAL
                                                                     ENTRY
                                                                                 SESSION
FULL ESTIMATED COST
                                                                      2.83
                                                                                  256.54
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
                                                               SINCE FILE
                                                                                   TOTAL
                                                                     ENTRY
                                                                                 SESSION
CA SUBSCRIBER PRICE
                                                                     -0.65
                                                                                   -0.65
```

STN INTERNATIONAL LOGOFF AT 14:56:43 ON 30 JAN 2003

LOGINID:ssspta1202txn

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

```
Welcome to STN International
                 Web Page URLs for STN Seminar Schedule - N. America
NEWS
                 "Ask CAS" for self-help around the clock
NEWS
         Apr 08
                 BEILSTEIN: Reload and Implementation of a New Subject Area
NEWS
         Apr 09
NEWS
         Apr 09
                 ZDB will be removed from STN
NEWS
         Apr 19
                 US Patent Applications available in IFICDB, IFIPAT, and IFIUDB
NEWS
        Apr 22
                 Records from IP.com available in CAPLUS, HCAPLUS, and ZCAPLUS
NEWS
        Apr 22
                BIOSIS Gene Names now available in TOXCENTER
NEWS
     8 Apr 22
                 Federal Research in Progress (FEDRIP) now available
     9 Jun 03
                New e-mail delivery for search results now available
NEWS
NEWS 10 Jun 10
NEWS 11 Jun 10
                 MEDLINE Reload
                 PCTFULL has been reloaded
NEWS 12 Jul 02
                 FOREGE no longer contains STANDARDS file segment
NEWS 13 Jul 22
                USAN to be reloaded July 28, 2002;
                 saved answer sets no longer valid
NEWS 14 Jul 29
                 Enhanced polymer searching in REGISTRY
NEWS 15 Jul 30
                NETFIRST to be removed from STN
NEWS 16 Aug 08
                 CANCERLIT reload
NEWS 17 Aug 08
                 PHARMAMarketLetter (PHARMAML) - new on STN
NEWS 18 Aug 08 NTIS has been reloaded and enhanced
NEWS 19 Aug 19 Aquatic Toxicity Information Retrieval (AQUIRE)
                 now available on STN
                 IFIPAT, IFICDB, and IFIUDB have been reloaded
NEWS 20 Aug 19
NEWS 21 Aug 19
                 The MEDLINE file segment of TOXCENTER has been reloaded
NEWS 22 Aug 26
                 Sequence searching in REGISTRY enhanced
NEWS 23 Sep 03
                 JAPIO has been reloaded and enhanced
NEWS 24 Sep 16 Experimental properties added to the REGISTRY file
NEWS 25 Sep 16 CA Section Thesaurus available in CAPLUS and CA
NEWS 26 Oct 01 CASREACT Enriched with Reactions from 1907 to 1985
NEWS 27 Oct 21 EVENTLINE has been reloaded
NEWS 28 Oct 24 BEILSTEIN adds new search fields
NEWS 29 Oct 24 Nutraceuticals International (NUTRACEUT) now available on STN
NEWS 30 Oct 25 MEDLINE SDI run of October 8, 2002
NEWS 31 Nov 18 DKILIT has been renamed APOLLIT
NEWS 32 Nov 25 More calculated properties added to REGISTRY
NEWS 33 Dec 02 TIBKAT will be removed from STN
NEWS 34 Dec 04 CSA files on STN
NEWS 35 Dec 17 PCTFULL now covers WP/PCT Applications from 1978 to date
NEWS 36 Dec 17 TOXCENTER enhanced with additional content
NEWS 37 Dec 17 Adis Clinical Trials Insight now available on STN
NEWS 38 Dec 30 ISMEC no longer available
NEWS 39 Jan 13 Indexing added to some pre-1967 records in CA/CAPLUS
NEWS 40 Jan 21 NUTRACEUT offering one free connect hour in February 2003
NEWS 41 Jan 21 PHARMAML offering one free connect hour in February 2003
NEWS 42 Jan 29
                 Simultaneous left and right truncation added to COMPENDEX,
                 ENERGY, INSPEC
NEWS EXPRESS January 6 CURRENT WINDOWS VERSION IS V6.01a,
              CURRENT MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP),
              AND CURRENT DISCOVER FILE IS DATED 01 OCTOBER 2002
NEWS HOURS
              STN Operating Hours Plus Help Desk Availability
NEWS INTER
              General Internet Information
NEWS LOGIN
              Welcome Banner and News Items
NEWS PHONE
              Direct Dial and Telecommunication Network Access to STN
NEWS WWW
              CAS World Wide Web Site (general information)
```

Enter NEWS followed by the item number or name to see news on that specific topic.

All use of STN is subject to the provisions of the STN Customer agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may result in loss of user privileges and other penalties.

FILE 'HOME' ENTERED AT 15:04:45 ON 30 JAN 2003

=> file reg COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 15:05:05 ON 30 JAN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 28 JAN 2003 HIGHEST RN 482573-45-5 DICTIONARY FILE UPDATES: 28 JAN 2003 HIGHEST RN 482573-45-5

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

=> Uploading 09910702.str

L1 STRUCTURE UPLOADED

=> d l1 L1 HAS NO ANSWERS L1 ST

property

Structure attributes must be viewed using STN Express query preparation.

=> s l1 SAMPLE SEARCH INITIATED 15:05:28 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED - 59715 TO ITERATE

1000 ITERATIONS 1.7% PROCESSED INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **INCOMPLETE** BATCH **INCOMPLETE**

PROJECTED ITERATIONS:

EXCEEDS 1000000

PROJECTED ANSWERS:

EXCEEDS

L2

0 SEA SSS SAM L1

=> s l1 ful

FULL SEARCH INITIATED 15:05:34 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED - >1,000,000 TO ITERATE

< 32.6% PROCESSED 389570 ITERATIONS (1 INCOMPLETE) 4 ANSWERS

< 33.5% PROCESSED 400000 ITERATIONS (1 INCOMPLETE) 4 ANSWERS

INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)

SEARCH TIME: 00.00.39

FULL FILE PROJECTIONS: ONLINE **INCOMPLETE**

BATCH **INCOMPLETE**

PROJECTED ITERATIONS:

EXCEEDS 1000000

PROJECTED ANSWERS:

EXCEEDS

4 SEA SSS FUL L1

=> file caplus

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST

148.55 148.76

O ANSWERS

FILE 'CAPLUS' ENTERED AT 15:06:32 ON 30 JAN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 30 Jan 2003 VOL 138 ISS 5 FILE LAST UPDATED: 29 Jan 2003 (20030129/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

```
09/ 910,702
```

=> s 13

T.4 2 L3

=> d l4 1- ibib abs hitstr

YOU HAVE REQUESTED DATA FROM 2 ANSWERS - CONTINUE? Y/(N):y

ANSWER 1 OF 2 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER:

2002:670837 CAPLUS

TITLE:

Supramolecular organization of oligopeptides, through

complexation with surfactants

AUTHOR (S):

General, Sascha; Antonietti, Markus

CORPORATE SOURCE:

Max Planck Institute of Colloids and Interfaces,

Potsdam-Golm, 14424, Germany

SOURCE:

Angewandte Chemie, International Edition (2002),

41(16), 2957-2960

CODEN: ACIEF5; ISSN: 1433-7851

PUBLISHER:

Wiley-VCH Verlag GmbH

DOCUMENT TYPE:

Journal

LANGUAGE:

English

Oligopeptides with a small no. of charged sites can be pptd. from water by complexation with oppositely charged surfactants, as exemplified here with oxidized glutathione, GSSG. These complexes are well-defined 1:1 species, dissolve in org. solvents, and form highly organized supramol. aggregates (soln.) or mesophases (solid-state films). This ionic self-assembly with surfactants represents a simple access to new peptide superstructures with structural features on the nanometer scale.

IT 480436-78-0

RL: PRP (Properties)

(prepn. and properties of oligopeptide-surfactant superassemblies)

480436-78-0 CAPLUS RN

Glycine, L-.gamma.-glutamyl-L-cysteinyl-, bimol. (2.fwdarw.2')-disulfide, CN mixt. with soya lecithins (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT:

22 THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 2 OF 2 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER:

2001:526075 CAPLUS

DOCUMENT NUMBER:

135:122506

TITLE:

Preparation of 2-amino-2-(aryl or heteroaryl)propanoic acid derivatives and related compounds as non-peptidyl inhibitors of VLA-4 dependent cell binding useful in treating inflammatory, autoimmune, and respiratory

diseases

INVENTOR(S):

Chupak, Louis Stanley; Duplantier, Allen Jacob; Lau,

Wan Fang; Milici, Anthony John

PATENT ASSIGNEE(S): SOURCE:

Pfizer Products Inc., USA

PCT Int. Appl., 182 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT	NO.		KI	ND :	DATE			A.	PPLI	CATI	ON NO	o. :	DATE			
								-								
WO 2001051487			A	1 20010719				WO 2000-IB1893 20001215								
. W:	ΑE,	AG,	AL,	AM,	ΑT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,
															GM,	
	HU,	ID,	ΙĻ,	IN,	IS,	JP,	KΕ,	KG,	ΚP,	KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,
															RO,	
	SD,	SE,	SG,	SI,	SK,	SL,	TJ,	TM,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VN,

```
YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                         BR 2000-16818
                                                               20001215
     BR 2000016818
                             20021001
                        Α
     EP 1244656
                             20021002
                                            EP 2000-983429
                        Α1
                                                                20001215
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
                                             US 2000-747246
     US 2002049236
                       A1
                             20020425
                                                                20001221
     US 2003004196
                        Α1
                             20030102
                                             US 2002-170289
                                                                20020612
                                             NO 2002-3085
     NO 2002003085
                             20020626
                                                                20020626
PRIORITY APPLN. INFO.:
                                          US 1999-173260P P 19991228
                                                            W
                                          WO 2000-IB1893
                                                               20001215
                                          US 2000-747246
                                                            B3 20001221
```

OTHER SOURCE(S): MARPAT 135:122506

$$R^4$$
|
N
B
CCH₂) n-Y
 R^2
 R^3
|
CCH₂) m-(CR⁷R⁸) p-CO₂H

There is disclosed a genus of non-peptidyl compds. represented by formula AB A-(CH2)n-Y-N(R4)-CR2R3-B-E-(CH2)m-(CR7R8)p-CO2H [A is (un)substituted C1-C6 alkyl, cycloalkyl, aryl, heteroaryl or heterocyclyl, A1-NHCONH-A2, A1-NHCO2-A2, A1-O2CNH-A2, A1-NHSO2NH-A2, A1-NHCO-A2, A1-CONH-A2, A-NHSO2-A2, etc. (where A1, A2 = H, (un) substituted aryl, C1-6 alkyl, C2-6 alkenyl, C2-6 alkynyl, cycloalkyl, heteroaryl, or heterocyclyl); E = a single bond, O, (un) substituted NH, CH:CH, C.tplbond.C, S, SO, SO2, (un) substituted CH2NH or CH2; B = Q-Q8 (proviso provided), etc. (where X =O, CO, S, SO, SO2, optionally substituted NH; X1, X2, X3 = optionally substituted CH, N; Y = a single bond, CO, CS, SO2); m = 0.1; n = 0.2; R2, R3 = H, (un)substituted C1-6 alkyl, C2-6 alkenyl, C3-14 carbocyclyl, heterocyclyl, C1-6 alkyl-OR5, C1-6 alkyl-SR5, C1-6 alkyl-SO2R5, heteroaryl, or aryl (where R5, R6 = H, optionally substituted C1-6 alkyl, C2-6 alkenyl, C2-6 alkynyl, aryl, cycloalkyl, heteroaryl, or heterocyclyl, CF3); R4 = H, (un)substituted C1-6 alkyl; R7 = C1-6 alkyl, (CH2)kOR5, (CH2) kCOR5, (CH2) kCONR6R5, (CH2) kNR6COR5, (CH2) k CO2 R5, (CH2) kNR6SO2R5, (CH2) kNR6R5, F, CF3, etc.; R8 = H, cyano, C1-6 alkyl or alkoxy]. These compds. are active as potent inhibitors of the binding of very late antigen-4 (VLA-4) to proteins such as vascular cell adhesion mol.-1 (VCAM-1), the HepII/IIICS domain (CS-1 region) of fibronectin and osteopontin (no data). They are effective for preventing, inhibiting, suppressing or reducing cell adhesion and consequent or assocd. pathogenic processes subsequently mediated by VLA-4. They are useful in treating inflammatory, autoimmune, and respiratory diseases which are selected from asthma, multiple sclerosis, rheumatoid arthritis, osteoarthritis, inflammatory bowel disease, psoriasis, host rejection following organ transplantation, atherosclerosis, and other diseases mediated by or assocd. with VLA-4. Thus, 3,5-dichlorobenzenesulfonyl chloride (86.7 mg) was added to a soln. of 2-allyloxycarbonylamino-3-(3-pyrrolidin-2ylisoxazol-5-yl)propionic acid Et ester hydrochloride (110 mg) and sodium carbonate (93.5 mg) in water (1.5 mL) and stirred overnight to give 37% 2-Allyloxycarbonylamino-3-[3-[1-(3,5-dichlorobenzenesulfonyl)pyrrolidin-2y]isoxazol-5-yl]propionic acid Et ester which (59 mg) was stirred with 2 M aq. LiOH (0.5 mL) at room temp. for 40 min and acidified to pH 1 with 1 M $\,$ HCl t give 91% 2-Allyloxycarbonylamino-3-[3-[1-(3,5dichlorobenzenesulfonyl)pyrrolidin-2-y]isoxazol-5-yl]propionic acid.

350675-19-3P 350675-20-6P 350675-21-7P

CN

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of amino(aryl or heteroaryl)propanoic acid derivs. and related compds. as non-peptidyl inhibitors of VLA-4 dependent cell binding for treating inflammatory, autoimmune, and respiratory diseases)

RN 350675-19-3 CAPLUS

5-Isoxazolepropanoic acid, 3-[1-[[4-(phenylmethoxy)phenyl]acetyl]-2-pyrrolidinyl]-.alpha.-(2-pyridinylamino)- (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 2-A

RN 350675-20-6 CAPLUS

CN 5-Isoxazolepropanoic acid, 3-[1-[[4-(phenylmethoxy)phenyl]acetyl]-2-pyrrolidinyl]-.alpha.-(3-pyridinylamino)- (9CI) (CA INDEX NAME)

PAGE 2-A

RN 350675-21-7 CAPLUS

CN

5-Isoxazolepropanoic acid, 3-[1-[[4-(phenylmethoxy)phenyl]acetyl]-2-pyrrolidinyl]-.alpha.-(4-pyridinylamino)- (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 2-A



REFERENCE COUNT:

10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> d his

(FILE 'HOME' ENTERED AT 15:04:45 ON 30 JAN 2003)

FILE 'REGISTRY' ENTERED AT 15:05:05 ON 30 JAN 2003

L1 STRUCTURE UPLOADED

L2 0 S L1

L3 4 S L1 FUL

FILE 'CAPLUS' ENTERED AT 15:06:32 ON 30 JAN 2003

L4 2 S L3

=> log y

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	7.67	156.43
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE	TOTAL
CA SUBSCRIBER PRICE	ENTRY -1.30	SESSION -1.30

STN INTERNATIONAL LOGOFF AT 15:07:08 ON 30 JAN 2003

Welcome to STN International! Enter x:x

```
LOGINID:ssspta1202txn
PASSWORD:
TERMINAL (ENTER 1, 2, 3, OR ?):2
 * * * * * * * *
                     Welcome to STN International
                 Web Page URLs for STN Seminar Schedule - N. America
NEWS
NEWS
         Apr 08
                 "Ask CAS" for self-help around the clock
NEWS
         Apr 09
                 BEILSTEIN: Reload and Implementation of a New Subject Area
NEWS
         Apr 09
                 ZDB will be removed from STN
NEWS
         Apr 19
                 US Patent Applications available in IFICDB, IFIPAT, and IFIUDB
NEWS
         Apr 22
                 Records from IP.com available in CAPLUS, HCAPLUS, and ZCAPLUS
         Apr 22
                 BIOSIS Gene Names now available in TOXCENTER
NEWS
      7
         Apr 22
                 Federal Research in Progress (FEDRIP) now available
NEWS
         Jun 03
                 New e-mail delivery for search results now available
NEWS
      9
NEWS 10
         Jun 10
                 MEDLINE Reload
         Jun 10
                 PCTFULL has been reloaded
NEWS 11
         Jul 02
                 FOREGE no longer contains STANDARDS file segment
NEWS 12
         Jul 22
                 USAN to be reloaded July 28, 2002;
 NEWS 13
                  saved answer sets no longer valid
NEWS 14
         Jul 29
                 Enhanced polymer searching in REGISTRY
NEWS 15
         Jul 30
                 NETFIRST to be removed from STN
NEWS 16 Aug 08
                 CANCERLIT reload
NEWS 17 Aug 08
                 PHARMAMarketLetter(PHARMAML) - new on STN
NEWS 18 Aug 08
                 NTIS has been reloaded and enhanced
NEWS 19
         Aug 19
                Aquatic Toxicity Information Retrieval (AQUIRE)
                 now available on STN
NEWS 20
         Aug 19
                 IFIPAT, IFICDB, and IFIUDB have been reloaded
NEWS 21 Aug 19
                 The MEDLINE file segment of TOXCENTER has been reloaded
NEWS 22 Aug 26
                 Sequence searching in REGISTRY enhanced
NEWS 23
         Sep 03
                 JAPIO has been reloaded and enhanced
NEWS 24 Sep 16 Experimental properties added to the REGISTRY file
NEWS 25 Sep 16 CA Section Thesaurus available in CAPLUS and CA
NEWS 26 Oct 01 CASREACT Enriched with Reactions from 1907 to 1985
NEWS 27 Oct 21 EVENTLINE has been reloaded
NEWS 28 Oct 24 BEILSTEIN adds new search fields
NEWS 29 Oct 24 Nutraceuticals International (NUTRACEUT) now available on STN
NEWS 30 Oct 25 MEDLINE SDI run of October 8, 2002
NEWS 31 Nov 18 DKILIT has been renamed APOLLIT
NEWS 32 Nov 25 More calculated properties added to REGISTRY
NEWS 33 Dec 02 TIBKAT will be removed from STN
NEWS 34 Dec 04 CSA files on STN
NEWS 35 Dec 17 PCTFULL now covers WP/PCT Applications from 1978 to date
NEWS 36 Dec 17 TOXCENTER enhanced with additional content
NEWS 37 Dec 17 Adis Clinical Trials Insight now available on STN
NEWS 38 Dec 30 ISMEC no longer available
NEWS 39 Jan 13 Indexing added to some pre-1967 records in CA/CAPLUS
NEWS 40 Jan 21 NUTRACEUT offering one free connect hour in February 2003
NEWS 41 Jan 21
                PHARMAML offering one free connect hour in February 2003
```

NEWS 42 Jan 29 Simultaneous left and right truncation added to COMPENDEX,

ENERGY, INSPEC

NEWS EXPRESS January 6 CURRENT WINDOWS VERSION IS V6.01a,

CURRENT MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP),

AND CURRENT DISCOVER FILE IS DATED 01 OCTOBER 2002

NEWS HOURS STN Operating Hours Plus Help Desk Availability

NEWS INTER General Internet Information NEWS LOGIN Welcome Banner and News Items

NEWS PHONE Direct Dial and Telecommunication Network Access to STN

NEWS WWW CAS World Wide Web Site (general information)

Enter NEWS followed by the item number or name to see news on that specific topic.

All use of STN is subject to the provisions of the STN Customer agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may result in loss of user privileges and other penalties.

FILE 'HOME' ENTERED AT 15:11:42 ON 30 JAN 2003

=> file reg COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST

0.21 SESSION 0.21

FILE 'REGISTRY' ENTERED AT 15:11:52 ON 30 JAN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 28 JAN 2003 HIGHEST RN 482573-45-5 DICTIONARY FILE UPDATES: 28 JAN 2003 HIGHEST RN 482573-45-5

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

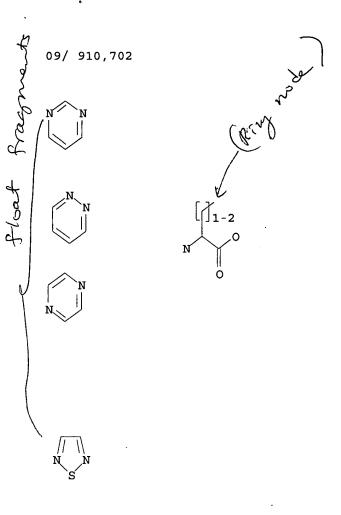
Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

Uploading 09910702.str

L1 STRUCTURE UPLOADED

=> d l1 L1 HAS NO ANSWERS L1 STR



Structure attributes must be viewed using STN Express query preparation.

=> s l1

SAMPLE SEARCH INITIATED 15:12:14 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED - 0 TO ITERATE

100.0% PROCESSED 0 ITERATIONS 0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

0 TO 0 0 TO 0 PROJECTED ITERATIONS:

PROJECTED ANSWERS:

L20 SEA SSS SAM L1

=> s l1 ful

FULL SEARCH INITIATED 15:12:19 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED - 1 TO ITERATE

100.0% PROCESSED 1 ITERATIONS 0 ANSWERS

SEARCH TIME: 00.00.01

L3 0 SEA SSS FUL L1

=> file marpat

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST 148.55 148.76

FILE 'MARPAT' ENTERED AT 15:12:55 ON 30 JAN 2003

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2003 American Chemical Society (ACC)

COPYRIGHT (C) 2003 American Chemical Society (ACS)

FILE CONTENT: 1988-PRESENT (VOL 104 ISS 15-VOL 138 ISS 4) (20030124/ED)

MOST RECENT CITATIONS FOR PATENTS FROM FIVE MAJOR ISSUING AGENCIES (COVERAGE TO THESE DATES IS NOT COMPLETE):

US 6500220 31 DEC 2002 DE 20211496 19 NOV 2002 EP 1269992 01 JAN 2003 JP 2002373833 26 DEC 2002 WO 2003000260 03 JAN 2003

Structure search limits have been raised. See HELP SLIMIT for the new, higher limits.

=> d his

(FILE 'HOME' ENTERED AT 15:11:42 ON 30 JAN 2003)

FILE 'REGISTRY' ENTERED AT 15:11:52 ON 30 JAN 2003

L1 STRUCTURE UPLOADED

L2 0 S L1 L3 0 S L1 FUL

FILE 'MARPAT' ENTERED AT 15:12:55 ON 30 JAN 2003

=> s 11

SAMPLE SEARCH INITIATED 15:13:07 FILE 'MARPAT'
SAMPLE SCREEN SEARCH COMPLETED - 32 TO ITERATE

93.8% PROCESSED 30 ITERATIONS 0 ANSWERS

100.0% PROCESSED 32 ITERATIONS (1 INCOMPLETE) 1 ANSWERS

SEARCH TIME: 00.00.19

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**

PROJECTED ITERATIONS: 301 TO 979
PROJECTED ANSWERS: 1 TO 80

L4 1 SEA SSS SAM L1

=> s l1 ful

FULL SEARCH INITIATED 15:13:30 FILE 'MARPAT'
FULL SCREEN SEARCH COMPLETED - 700 TO ITERATE

93.1% PROCESSED 652 ITERATIONS (2 INCOMPLETE) 2 ANSWERS

96.3% PROCESSED 674 ITERATIONS (3 INCOMPLETE) 3 ANSWERS

100.0% PROCESSED 700 ITERATIONS (7 INCOMPLETE) 7 ANSWERS

SEARCH TIME: 00.00.48

L5 7 SEA SSS FUL L1

=> file caplus

COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
105.35
254.11

FILE 'CAPLUS' ENTERED AT 15:14:36 ON 30 JAN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 30 Jan 2003 VOL 138 ISS 5 FILE LAST UPDATED: 29 Jan 2003 (20030129/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 15L6 7 L5

=> d l6 1- ibib abs hitstr YOU HAVE REQUESTED DATA FROM 7 ANSWERS - CONTINUE? Y/(N):y

ANSWER 1 OF 7 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 2002:90026 CAPLUS

DOCUMENT NUMBER:

136:135019

TITLE:

Preparation of 3-amino-2-(4-aminocarbonyloxy)phenylpropionic acid derivatives as antiinflammatory agents

and .alpha.4 Integrin inhibitors

INVENTOR(S):

Konradi, Andrei W.; Pleiss, Michael A.; Thorsett, Eugene D.; Ashwell, Susan; Welmaker, Gregory S.;

Kreft, Anthony; Sarantakis, Dimitrios; Dressen, Darren

B.; Grant, Francine S.; Xu, Ying-Zi

PATENT ASSIGNEE(S):

Elan Pharmaceuticals, Inc., USA; American Home

Products Corporation PCT Int. Appl., 137 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

SOURCE:

English

FAMILY ACC. NUM. COUNT:

```
PATENT NO.
                  KIND DATE
                                      APPLICATION NO. DATE
    -----
                         _____
                                       ------
    WO 2002008206
                   A1 20020131
                                      WO 2001-US23073 20010720
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT,
            RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ,
            VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
            DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
            BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
    US 2002055509
                   A1 20020509
                                      US 2001-910685 20010720
PRIORITY APPLN. INFO.:
                                     US 2000-220134P P 20000721
OTHER SOURCE(S):
                      MARPAT 136:135019
```

$$R^4$$
 (Alk) n CR (R³) CH₂N (R?) Ar R^5 OCONR¹R²

AB 3-Amino-2-(4-aminocarbonyloxy)phenyl-propionic acid derivs. I wherein R is a carboxylic acid; R1 and R2 are independently selected from the group consisting of hydrogen, alkyl, substituted alkyl, alkenyl, substituted alkenyl, cycloalkyl, substituted cycloalkyl, or R1 and R2, together with the nitrogen atom to which they are attached, are joined to form an optionally substituted heterocyclic ring provided that said substituted alkyl, substituted alkenyl and substituted cycloalkyl do not carry an aryl, substituted aryl, heteroaryl or substituted heteroaryl group; Ra and R3 are independently a hydrogen or a Me group; R4 and R5 are independently selected from the group consisting of heteroatom group; n is zero or an integer 1; Alk is a straight or branched alkylene chain; Ar is an optionally substituted arom. or heteroarom. group, as well as their pharmaceutical use as .alpha.4.beta.7 Integrin inhibitors for the treatment of inflammatory diseases. Thus, 3-[4-(3,5-dichloropyrid-4ylcarboxamido)phenyl]-2-(3-chlorophenylamino)propanoic acid was prepd. as .alpha.4 Integrin inhibitor. The preferred compds. of the invention generally have IC50 values in the .alpha.4.beta.1 and .alpha.a.beta.7 assays of 1 .mu.M and below. In the other assays featuring .alpha. integrins of other subgroups the same compds. had IC50 values of 50 .mu.M and above thus demonstrating the potency and selectivity of their action against .alpha.4 integrins. Title compds. were prepd. for treating an inflammatory condition in a mammalian patient which condition is mediated by Very Late Antigen 4 (VLA-4). Inflammatory condition is selected from the group consisting of asthma, Alzheimer's disease, atherosclerosis, AIDS dementia, diabetes, inflammatory bowel disease, multiple sclerosis, rheumatoid arthritis, tissue transplantation, tumor metastasis, meningitis, encephalitis, stroke, nephritis, retinitis, atopic dermatitis, psoriasis, myocardial ischemia and acute leukocyte-mediated lung injury. THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 2

Ι

ANSWER 2 OF 7 CAPLUS COPYRIGHT 2003 ACS L6 2002:90023 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 136:135018

Preparation of 3-(heteroaryl) alanine derivatives as TITLE: inhibitors of leukocyte adhesion mediated by VLA-4

INVENTOR(S): Konradi, Andrei W.; Pleiss, Michael A.; Thorsett,

Eugene D.; Ashwell, Susan; Welmaker, Gregory S.; Kreft, Anthony; Sarantakis, Dimitrios; Dressen, Darren

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

B.; Grant, Francine S.; Semko, Christopher; Xu,

Ying-Zi; Stappenbeck, Frank

PATENT ASSIGNEE(S): Elan Pharmaceuticals, Inc., USA; American Home

> Products Corporation PCT Int. Appl., 132 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT:

```
APPLICATION NO. DATE
                                       KIND DATE
         PATENT NO.
                                                  _____
                                                                              -----
                                       ----
                                                                              WO 2001-US23097 20010720
         WO 2002008203
                                         A2
                                                  20020131
         WO 2002008203
                                        A3
                                                  20020523
                W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
                       CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
               CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, F1, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                                  20020502
                                                                             US 2001-910466 20010719
         US 2002052375
                                        A1
PRIORITY APPLN. INFO.:
                                                                         US 2000-220131P P 20000721
                                             MARPAT 136:135018
OTHER SOURCE(S):
```

3-(Heteroaryl) alanine derivs. I [A = an (un) substituted aryl, heteroaryl, AB cycloalkyl, or heterocyclic group; R2 = a nitrogen contg. (un) substituted, heteroaryl; Y = (CH2)m; m = 0 or 1; R1 = H, (un)substituted, alkyl, alkenyl, cycloalkyl, cycloalkenyl, aryl, heteroaryl, or heterocyclic; X = OH, (un) substituted alkoxy, alkenoxy, cycloalkoxy, cycloalkenoxy, aryloxy, heteroaryloxy, heterocyclyloxy, or NR3R3 [R3 = H, (un) substituted alkyl, alkenyl, cycloalkyl, aryl, heteroaryl, or heterocyclic]] were prepd. as inhibitors of leukocyte adhesion mediated by VLA-4. Compds. I have binding affinity to VLA-4 as expressed by an IC50 of about 15 .mu.M or less. Thus, N-[5-(2,2,2-trifluoroethyl)pyrimidin-4-yl]-DL-3-[5-(2,5dimethoxyphenyl)pyridin-2-yl]alanine was prepd. by multistep procedure via coupling of DL-[5-(2,6-dimethoxyphenyl)pyridine-2-yl]alanine Et ester and 4,6-dichloro-5-(2,2,2-trifluoroethyl)pyrimidine.

ANSWER 3 OF 7 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 1997:340728 CAPLUS

DOCUMENT NUMBER: 126:305589

Preparation of aryloxypyrimidines and related TITLE:

compounds as herbicides.

INVENTOR(S): Rheinheimer, Joachim; Vogelbacher, Uwe Josef; Baumann,

Ernst; Mislitz, Ulf; Westphalen, Karl-Otto; Walter,

PATENT ASSIGNEE(S): BASF A.-G., Germany SOURCE: Ger. Offen., 44 pp.

CODEN: GWXXBX

DOCUMENT TYPE: Patent German LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT NO.	KIND	DATE	APPLICATION NO. D	DATE
DE 19536809	A1	19970403	DE 1995-19536809 1	9951002
WO 9712879	A1	19970410	WO 1996-EP4204 1	19960926
W· CA CN	TP KR	IIS		

RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE EP 873318 19981028 EP 1996-932599 19960926 **A**1

R: AT, BE, CH, DE, ES, FR, GB, IT, LI, NL

CN 1996-198318 19960926 19981216 CN 1202158 Α JP 2000500122 ΤЭ 20000111 JP 1997-513945 19960926 PRIORITY APPLN. INFO.: DE 1995-19536809 19951002

WO 1996-EP4204 19960926

OTHER SOURCE(S): MARPAT 126:305589

GI

$$\begin{array}{c|c}
 & R^1 \\
 & N \\
 & N \\
 & N \\
 & R^2
\end{array}$$

Title compds. [I; A = substituted 5-membered heteroaryl; X = O, S; Y = N, AB CH; R1, R2 = halo, alkyl, haloalkyl, alkoxy, haloalkoxy, alkylthio, alkylamino, dialkylamino; R3 = H, (substituted) 5-membered heteroaryl, succinyliminooxy, etc.], were prepd. Thus, 2,2-dimethyl-5-(2methoxythiazol-5-yl)-4H-1,3-benzodioxin-5-one (prepn. given) was refluxed 4 h with NaOH and Bu4NOH in H2O and the product was stirred with KOCMe3 in Me2SO followed by addn. of 4,6-dimethoxy-2-methylsulfonylpyrimidine to qive 2-(2,6-dimethoxypyrimidin-2-yloxy)-6-(2-methylthiazol-5-yl)benzoic acid. The latter at 0.0156 kg/ha postemergent gave 100% control of Amaranthus retroflexus.

ANSWER 4 OF 7 CAPLUS COPYRIGHT 2003 ACS 1997:69790 CAPLUS

Ι

ACCESSION NUMBER:

DOCUMENT NUMBER: 126:89393

Preparation of diazathiabicycloalkanones and analogs TITLE:

as thrombin inhibitors

Dimaio, John; Gillard, John W.; Siddiqui, M. Arshad; INVENTOR (S):

Bachand, Benoit; Doherty, Annette Marian; Edmonds,

Jeremy John

Biochem Pharma Inc., Can.; Dimaio, John; Gillard, John PATENT ASSIGNEE(S):

W.; Siddiqui, M. Arshad; Bachand, Benoit; Doherty,

Annette Marian; Edmonds, Jeremy John

PCT Int. Appl., 87 pp. SOURCE:

CODEN: PIXXD2

Patent

DOCUMENT TYPE: English

LANGUAGE:

FAMILY ACC. NUM. COUNT: 1

PA	CENT	NO.		KI	ND :	D DATE APPLICATION NO. DATE											
									-	-							
WO 9637497 A1 19961:				1128 WO 1996-CA318 19960522													
	W:	AL,	AM,	ΑT,	AU,	ΑZ,	BB,	ВG,	BR,	BY,	CA,	CH,	CN,	CZ,	DE,	DK,	EE,
		ES,	FI,	GB,	GE,	HU,	IS,	JP,	KΕ,	KG,	ΚP,	KR,	KZ,	LK,	LR,	LS,	LT,
		LU,	LV,	MD,	MG,	MK,	MN,	MW,	MX,	NO,	NZ,	PL,	PT,	RO,	RU,	SD,	SE,
		SG,	SI														
	RW:	KΕ,	LS,	MW,	SD,	SZ,	UG,	ΑT,	BE,	CH,	DE,	DK,	ES,	FI,	FR,	GB,	GR,
		ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN	
CA	2218	816		A	A	1996	1128		C	A 19	96-2	2188	16	1996	0522		
AU	9656	825		Α	1	1996	1211		Αl	J 19	96-5	6825		1996	0522		
7.Δ	9604	090		Δ		1997	0513		7.7	4 19	96-4	090		1996	0522		

EP 846120 19980610 EP 1996-914817 19960522 **A1**

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,

IE, FI

19990518 JP 11505260 T2 JP 1996-535221 19960522

PRIORITY APPLN. INFO.: GB 1995-10264 19950522 Α W 19960522 WO 1996-CA318

MARPAT 126:89393 OTHER SOURCE(S):

GI

$$R^{3}N$$
 R^{7}
 R^{8}
 R^{2}
 R^{2}
 R^{2}
 R^{3}
 R^{2}
 R^{3}
 R^{4}
 R^{2}
 R^{2}
 R^{3}
 R^{4}
 R^{5}
 R^{6}
 R^{7}
 R^{8}
 R^{9}
 R^{1}
 R^{2}
 R^{3}
 R^{4}
 R^{2}
 R^{2}
 R^{3}
 R^{4}
 R^{5}
 R^{5}
 R^{6}
 R^{7}
 R^{8}
 R^{1}
 R^{2}
 R^{3}
 R^{4}
 R^{5}
 R^{5}
 R^{7}
 R^{7}
 R^{8}
 R^{1}
 R^{2}
 R^{3}
 R^{4}
 R^{5}
 R^{5}
 R^{5}
 R^{7}
 R^{7}
 R^{8}
 R^{1}
 R^{1}
 R^{2}
 R^{3}
 R^{4}
 R^{5}
 $R^{$

AΒ Title compds. [I; R2 = H, NH2, (ar)alkyl, etc.; R3,R4 = H, NR6R7, alkyl, aryl, etc.; R6 = polar amino acid residue, arginyl, etc.; R7,R8 = H or alkyl; Z = CHR5, O, SOO-2, etc.; R5 = H, alkyl, aryl, etc.; m,n = 0-2]were prepd. Thus, oxothiazolopyrimidinecarboxylate II (R = OH) (prepn. given) was amidated by HR1 (R1 = arginine residue Q, R9 = CO2CH2Ph) to give, after deprotection, diastereomers of II (R = Q, R9 = H). Data for in vitro biol. activity of I were given.

ANSWER 5 OF 7 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 1993:613908 CAPLUS

DOCUMENT NUMBER: 119:213908

TITLE: Silver halide photographic material

INVENTOR (S): Fukuwa, Junichi; Kobayashi, Akira; Goto, Kenji

PATENT ASSIGNEE(S): Konica Co., Japan

SOURCE: Can. Pat. Appl., 71 pp.

CODEN: CPXXEB

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
CA 2065106	AA	19921005	CA 1992-2065106	19920403
JP 05197057	A2	19930806	JP 1992-110787	19920403
PRIORITY APPLN. INFO.	:	JP	1991-99626	19910404
OTHER SOURCE(S):	MΔ	RPAT 119-213908		



A Ag halide photog. material for high-contrast dot image formation is disclosed. The material comprises a support and provided thereon a Aq halide emulsion layer and layers adjacent to the emulsion layer. The emulsion is subjected to desalinization comprising using denatured gelatin in the process of prepn. thereof. At least one of the layers contains a hydrazine deriv. and a compd. selected from the group consisting of those represented by formulas A(CH2)nSC(:N+HR1)NHR1 X- (A = OH, SO3-, or N(R2)2; R1 = H, (substituted) alkyl having 1-5 C atoms, or (substituted) Ph; R2 = (substituted) alkyl having 1-5 C atoms; X-= an anion), (R3)2N(CH2)nSC(S)N(R4)2 (R3 = H, (substituted) alkyl having 1-5 C atoms, or (substituted) aryl; R4 = (substituted) alkyl having 1-5 C atoms or (substituted) Ph; n = an integer of 2-5), or I (Q = a group of atoms necessary to form a 5- or 6-membered heterocyclic ring which may be condensed with a benzene or heterocyclic ring; M = H, an alkali metal atom, an ammonium group, or an amine residue).

ANSWER 6 OF 7 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 1993:560727 CAPLUS

DOCUMENT NUMBER:

119:160727 TITLE: Preparation of uronic acid (FR-900493) derivatives as

antibacterial agents

PATENT ASSIGNEE(S):

Fujisawa Pharmaceutical Co., Ltd., Japan

SOURCE:

Jpn. Kokai Tokkyo Koho, 57 pp.

CODEN: JKXXAF

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE ____ -----JP 05078385 A2 19930330 JP 1991-196172 19910225 PRIORITY APPLN. INFO.: GB 1990-4407 19900227 OTHER SOURCE(S): MARPAT 119:160727

GΙ

The title compds. [I; R1, R2 = (un)substituted NH2; R3 = (un)protected AΒ CO2H; R4 = H, OR9; R5 = H, OR10; R6 = OR11; R7 = H, R12; R9 - R12 = H, HO-protective group, or R9R10, R11R12 = (un)substituted lower alkylene; R8 = H, halo; X = N, N+R13; R13 = lower alkyl; Z = acid residue; n = 0,1; provided that when X = N+R13, R3 = protected CO2H and n = 1 or R3 = CO2H and n = 0] are prepd. Thus, 0.52 g di-tert-Bu dicarbonate and 25 mL H2O were added to a soln. of 1.04 g FR-900493 (II; X = X1 = H) in 15 mL 1,4-dioxane and then the mixt. was stirred at room temp. for 10 h to give, after column chromatog. using Diaion HP-20, II (R1 = H2N, R2 = Me3CO2CNH). II [R1 = 4-[Me(CH2)70]C6H4CH2CONH, R2 = H2N] showed min. inhibitory concn. of 6.25 and 12.5 .mu.g/mL against Staphylococcus aureus and Escherichia coli, resp. A total of 178 I including their salts were prepd.

Ι

ANSWER 7 OF 7 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER:

1993:560110 CAPLUS

DOCUMENT NUMBER:

119:160110

TITLE:

Thiourea derivatives and methods for inhibition of HIV

and related viruses

INVENTOR(S):

Lind, Peter Thomas; Morin, John Michael, Jr.; Noreen,

Rolf; Ternansky, Robert John

PATENT ASSIGNEE(S):

Medivir AB, Swed.

SOURCE:

PCT Int. Appl., 550 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

1

	KIND DATE	APPLICATION NO. DATE								
WO 9303022	A1 19930218	WO 1992-SE533 19920803								
W: AT, AU,	BB, BG, BR, CA, CH,	CS, DE, DK, ES, FI, GB, HU, JP, KP,								
KR, LK,	LU, MG, MN, MW, NL,	NO, PL, RO, RU, SD, SE								
RW: AT, BE,	CH, DE, DK, ES, FR,	GB, GR, IE, IT, LU, MC, NL, SE, BF,								
BJ, CF,	CG, CI, CM, GA, GN,	ML, MR, SN, TD, TG								
IL 102548	A1 19980816	IL 1992-102548 19920717								
AU 9220451	A1 19930311	AU 1992-20451 19920721								
AU 657978	B2 19950330									
CZ 282900	B6 19971112	CZ 1992-2288 19920722								
NO 9202949	A 19930203	NO 1992-2949 19920724								

ZA	9205663		Α	1994	0128		ZA	1992-	5663		1992	0728
ES	2051641		A:	1 1994	0616		ES	1992-	1601		1992	0729
ES	2051641		В:	1 1995	1001							
CN	1069882		Α	1993	0317		CN	1992-	11063	0	1992	0730
RU	2106341		C	1998	0310		RU	1992-	50527	83	1992	0730
CA	2075173		A.	A 1993	0203		CA	1992-2	20751	73	1992	0731
JP	0532013	8	A2	2 1993	1203		JΡ	1992-2	22521	8	1992	0731
AU	9224074		A.	1 1993	0302		AU	1992-2	24074		1992	0803
EP	540143		A	2 1993	0505		ΕP	1992-3	30709	2	1992	0803
EP	540143		A.	3 1995	0104							
	R: AT	, BE,	CH,	DE, DK,	FR,	GB, I	Ε,	IT, LI	, LU,	NL	, PT,	SE
NO	9301337		Α	1993	0203		NO	1993-	1337		1993	0407
ИО	9301338		Α	1993	0203		NO	1993-	1338		1993	0407
NO	9301339		Α	1993	0203		NO	1993-	1339		1993	0407
ИО	9301340		Α	1993	0203		NO	1993-	1340		1993	0407
US	5593993		Α	1997	0114		US	1995-3	39570	2	1995	0228
US	5658907		Α	1997	0819		US	1995-	45534	7	1995	0531
US	5714503		Α	1998	0203		US	1995-	45521	7	1995	0531
PRIORITY	APPLN.	INFO	.:			US	19:	91-739	927	Α	1991	0802
						NC	19:	92-294	9	A1	1992	0724
						US	19:	92-921	890	B2	1992	0729
						WC	19:	92-SE5	33	Α	1992	0803
	•					US	19	93-119	40	В3	1993	0201
						US	19	95-395	702	А3	1995	0228
OTHER SC	TIPCE (C)			ידעססעא	119.1	160110	١					

OTHER SOURCE(S):

MARPAT 119:160110

GΙ

AB A method for the inhibition of the replication of HIV is claimed that comprises contacting HIV with thiourea derivs. Among the specifically claimed compds. is N-[2-(2-pyridyl)ethyl]-N'-(5-bromo-2-pyridyl)thiourea (I).

=> d his

(FILE 'HOME' ENTERED AT 15:11:42 ON 30 JAN 2003)

I

FILE 'REGISTRY' ENTERED AT 15:11:52 ON 30 JAN 2003 L1 STRUCTURE UPLOADED

L2 0 S L1 L3 0 S L1 FUL

FILE 'MARPAT' ENTERED AT 15:12:55 ON 30 JAN 2003

L4 1 S L1 L5 7 S L1 FUL

FILE 'CAPLUS' ENTERED AT 15:14:36 ON 30 JAN 2003 L6 7 S L5

=> log y

COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
17.32
271.43

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
SINCE FILE TOTAL
ENTRY SESSION
CA SUBSCRIBER PRICE
-4.56

STN INTERNATIONAL LOGOFF AT 15:15:10 ON 30 JAN 2003